Inhibition of blood cholinesterase activity is a poor predictor of acetylcholinesterase inhibition in brain regions of guinea pigs exposed to repeated doses of low levels of soman.

Sally M. Anderson

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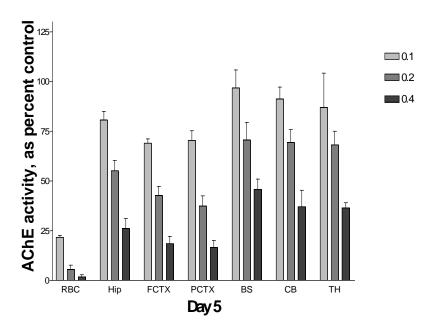
The experiment:

Expose male guinea pigs to subclinical doses of soman (0.1, 0.2, and 0.4 LD50) daily (M-F) for two weeks.

Collect blood and brain regions on exposure Day 5 and exposure Day 12, and after 3 and 17 days of recovery (Day 15 and Day 29).

Characterize inhibition of cholinesterase activity in blood and several brain regions: frontal cortex, parietal cortex, Hippocampus, cerebellum, brain stem and thalamus. There is great inhibition of AChE in red blood cells after 5 days of exposure to 0.1, 0.2, or 0.4 LD50 soman.

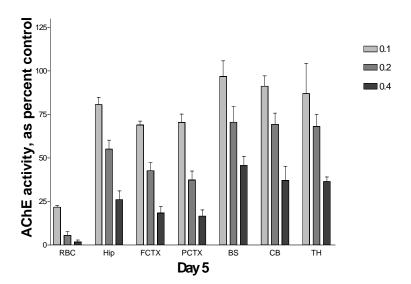
AChE inhibition in brain is significant but not as extreme As in red blood cells.

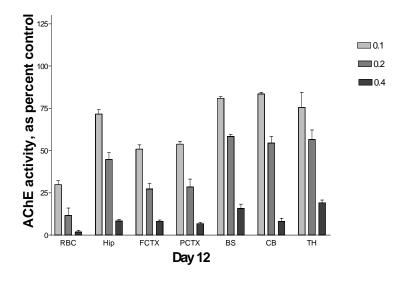


Soman inhibition of AChE in red blood cells is not greater on Day 12 than on Day 5.

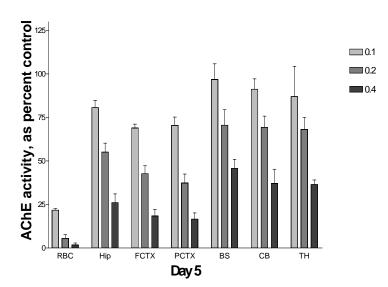
Inhibition of brain AChE by soman is greater in Day 12 Than Day 5.

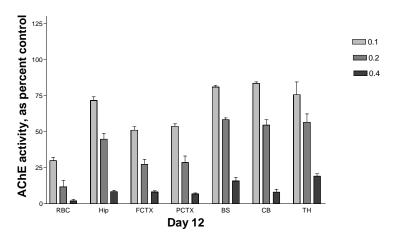
Inhibition of AChE in red blood cells by 0.1 or 0.2 LD50 Soman is much greater than inhibition in brain.

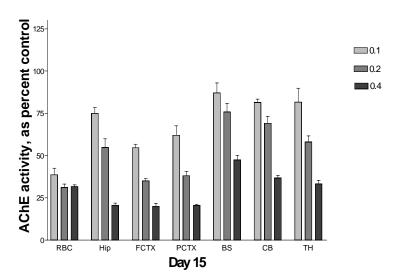




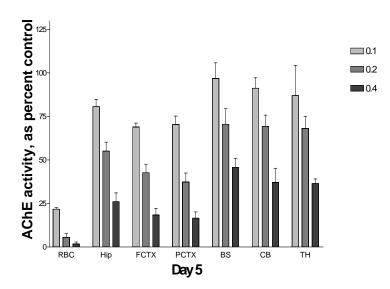
Recovery of AChE activity is apparent 3 days after the last exposure to soman.

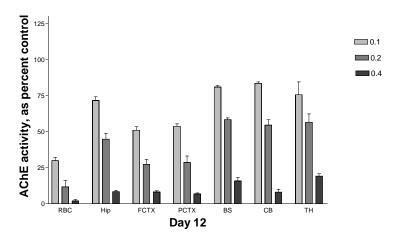


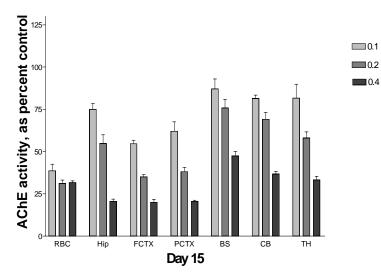


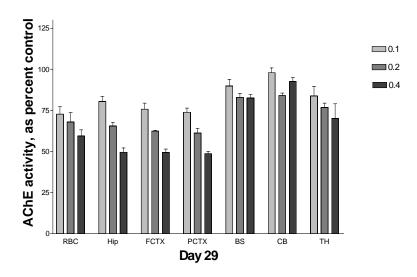


Seventeen days after exposure AChE activity is still less than control levels in red blood cells and several brain regions.

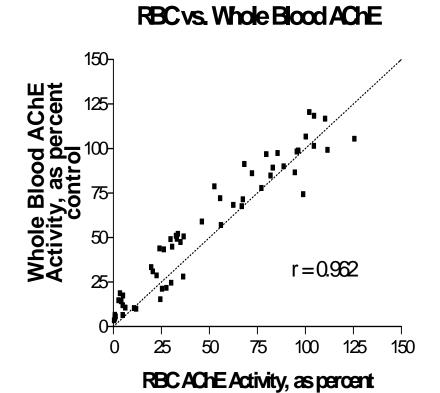




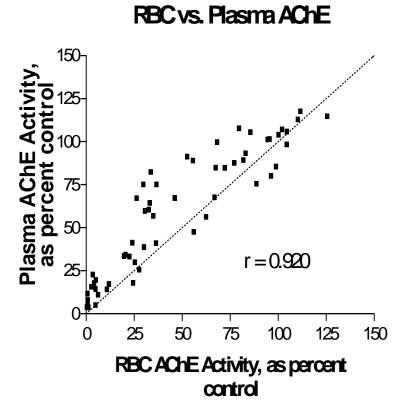




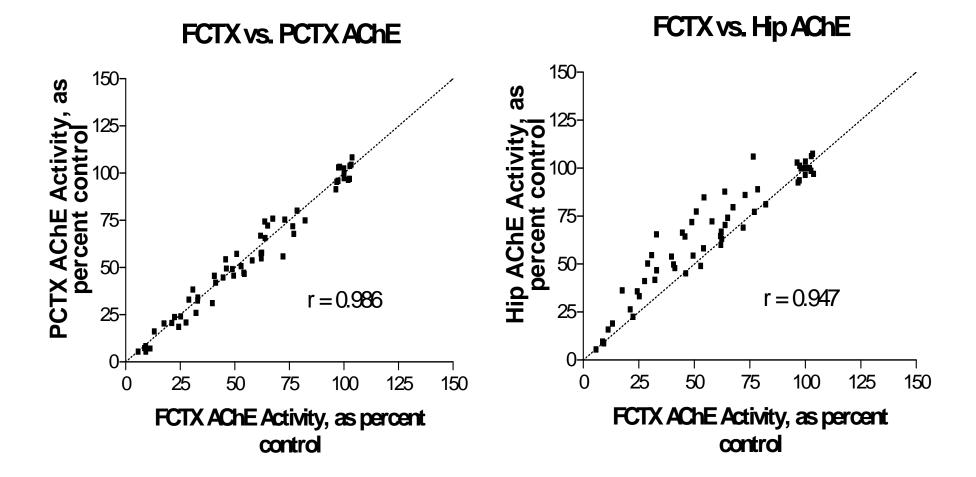
Amount of inhibition of AChE activity was highly correlated between whole blood, and red blood cells, and plasma.



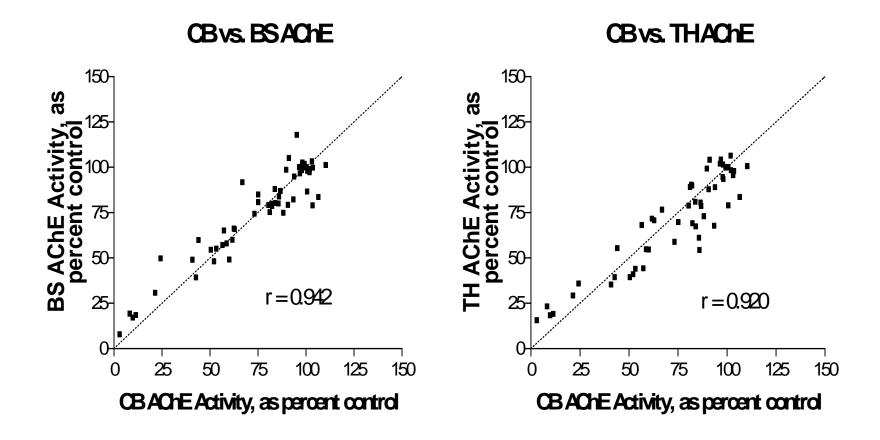
control



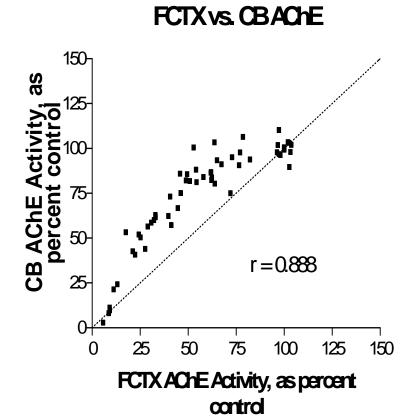
Amount of inhibition of AChE activity in frontal cortex is highly correlated with amount of inhibition in parietal cortex and in hippocampus.

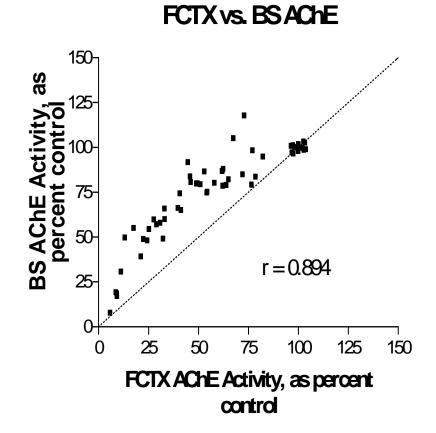


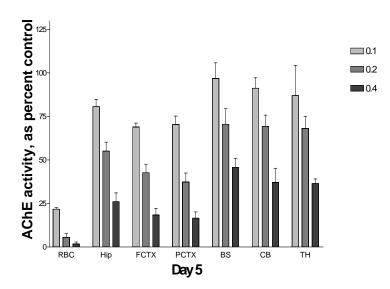
Amount of inhibition of AChE activity in cerebellum Is highly correlated with inhibition in brain stem and Thalamus.

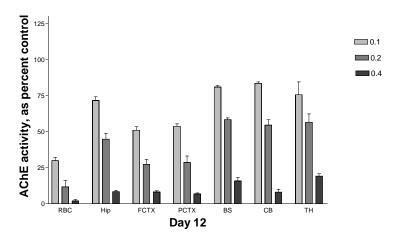


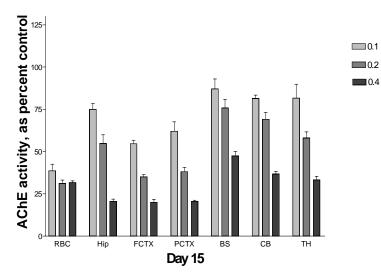
Amount of inhibition in brain stem, cerebellum, and thalamus is not as well correlated with inhibition in cortical regions or hippocampus.

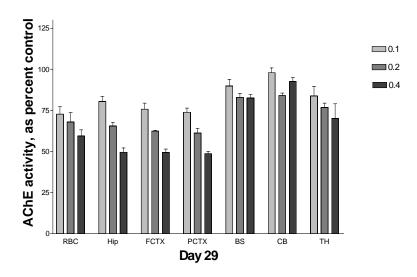






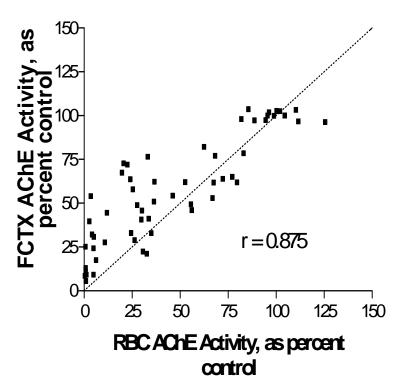




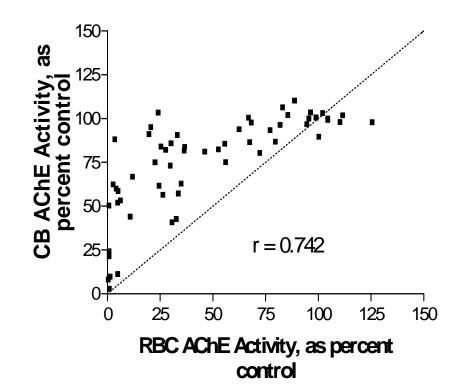


Correlations between inhibition of AChE in red blood cells amount of inhibition in brain was only modest.

RBCvs. FCTX AChE



RBC vs. CB AChE



	RBC AChE		
Frontal Cortex AChE	0.875		
Parietal Cortex AChE	0.871		
Hippocampus AChE	0.776		
Cerebellum AChE	0.742		
Brainstem AChE	0.723		
Thalamus AChE	0.788		

Inhibition of AChE in red blood cells is a poor predictor of AChE inhibition in brain regions of guinea pigs exposed to repeated doses of low levels of soman.

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